

Nern et al.

Appl. No.: 10/054,399 Atty Docket: DYOU13.1A2CP1

FIG. 1A

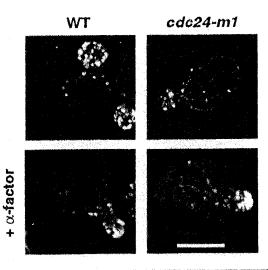


FIG. 1B

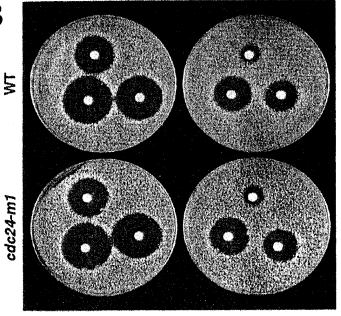
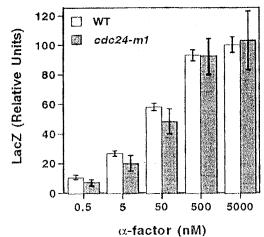


FIG. 1C



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LEOTIDE SEQUENCES AND PROTEIN

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FIG. 2A

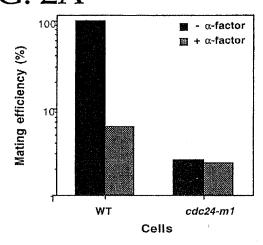
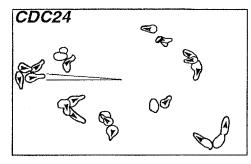


FIG. 2B



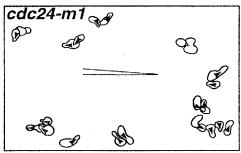
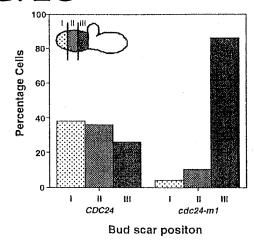
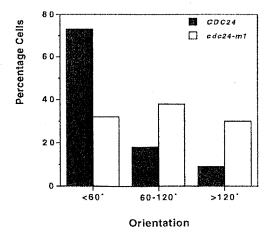


FIG. 2C





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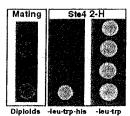
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FIG. 3A

FIG. 3B

AD STE4 CDC42 BEM1





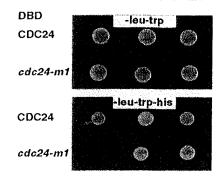


FIG. 3C

Cdc24

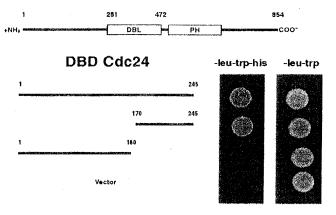
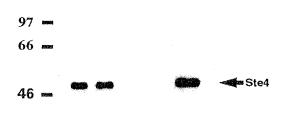


FIG. 3D







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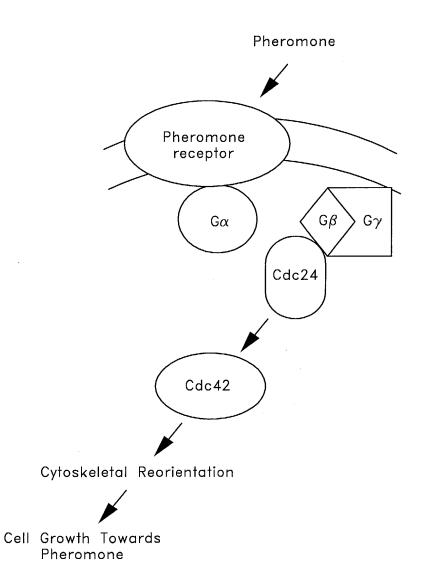


FIG. 4



LEOTIDE SEQUENCES AND PROTEIN SI Nern et al.

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FIG. 5_A

FIG. 5_B

FIG. 5

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LEOTIDE SEQUENCES AND PROTEIN SI

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5																			
	2113/1 ATG GAA	as m	G G.	CC3	aax	CCT	ama	ארשא	א כי א	2143		אככ	C N N	ጥሮል	λСТ	מירים	ጥረጥ	ጥጥር፤	ידיממ
	M E	H	P	P	A	A	L L	R	T		S		Q	S	T	s	s	L	N
	2173/21									2203	/31								
10	TCA GTA												GGC G	CCA P	GTC	AAT N	ATA I	AAC N	AAT N
	S V 2233/41	S	T	V	S	s	S	R	I	V 2263	S 1/51	S	G	P	V	14	±	IA	14
	TTC AAT	AAA	CCA	AGT	ACT	CCC	AAA	GAC	CAT	TTA	TTC	TAT	CGÁ	TGT	GAA	TCA	CTA	AAA	CGA
	F N	ĸ	P	s	T	P	K	D	H	L	F	Y	R	C	E	S	L	ĸ	R
15	2293/61 AAA CTA	~		* ma	a a m	000	N MCI	C 3 3	CON	2323	771	אארי	ርግ አ አ	COT	TO THE	አልጥ	CAG	CCM	CNA
	AAA CTA K L	CAA Q	AAA K	I	P	G	M	E	P		L	N	Q	A	F	N	O	A	E
	2353/81									2383	/91								
	CAA CTC															GGA	AAT		
20	Q L 2413/103	S	E	Q	Q	A	L	A	L		Q :/113	E	R	S	N	G	N	G	Н
	AGT AAT	GGC	AAA	CGT	CAT	CAA	TCA	TTA	GAC	GGT	GCC	ATG	AAT	AGA	CTT	TCA	GTT	GGT	TCT
	s N	G	K	R	H	Q	S	L	D	G	A	М	И	R	L	S	V	G	S
25	2473/123 GAT AGT	N COMP	Tr.CO	እጥሮ	מאם	CCT	ጥር አ	ጥጥር፤	ACA	2503	/ 131 ልጥር	L GCC	A CC	ΔΔΤ	GCG	TCA	ACG	TCA	TOT
23	D S	S	S	I	Q	G	S	L	T			A	T	N	A	s	T	s	s
	2533/141	L			_					2563	/153	L				~~~	~~~		~~*
	TTA ATC		GGT G	ATG M	CCA P	AAC N	AAC N	AAC N	ACT T		TTT	ACG	TTT	ACT T	GCA A	GGG	GTT V	L	P
30	L I 2593/161	s L	-							2623	/171	L	_			_	-	• • • • • • • • • • • • • • • • • • • •	_
	GCT AAT	ATT																	
	A N	Ι	S	V	D	P	A	T	H	L 2683	W :/191	K	L	F	Q	Q	G	A	P
	2653/181 TTT TGT	GTT	CTT	ATC	AAT	CAT	ATC	CTT	CCT	GAT	TCC	CAA	ATA	CCA	GTT	GTC	AGT	TCT	GAT
35	F C	V	L	I	N	H	I	L	Þ	D	S	Q	I	P	V	V	S	S	D
	2713/201 GAC TTG	אכיא	יוייני א	TCC	מממ	A A A	тсъ	GTA	ጥልጥ	2743	721] ጥጥጥ	ኒ ጥጥል	ידידא	GCC	GTC	AAG	ACA	CAA	TTG
	D L	R	I	Ĉ	K	ĸ	s	v	Y		F	L	I	A	v	K	T	Q	L
4.0	2773/221 AAT TTT						mm.a	3 CIM		2803	/231	C mm	mma	maa	C N C	* * *	aaa	(1 X X	CATI
40		GAT D	D	E	AAT N	M	F	T	I		MAT N	V	F	s	D	N	A	0	D
	2833/241		_							2863	/253	Ļ						_	,
	TTA ATC														TCA S	GAT D	GCT A	AGT S	GAC D
45	L I 2893/261	K	I	Ι	D	V	I	N	K	L 2923	上 [/27]	A	E	Y	5	D	A	3	D
73	CTG GGT	GGT	GGC	GAT	GAA					GAT	GTT	CAA							
	S G	G	G	D	E	D	V	N	M			Q	I	T	D	E	R	S	K
	2953/281 GTT TTC	CGA	GAA	ATT	ATC	GAA	ACA	GAA	AGA	2983 AAA	TAT	GTT	CAA	GAC	TTG	GAA	CTA	ATG	TGT
50	V F	R	E	I	I	E	T	E	R	ĸ	Y	V	Q	D	L	E	L .	M	С
	3013/301 AAA TAC		C1 70 70	C) A MI	CTDA	א מווימי	ממיי	acc	<i>ር</i> ግ አ አ	3043	7311 TTC	anton. T	ም ር አ	CAA	ממים	ידיידי מ	CAC	ጥጥር	מירידי
	K Y	R	O	D	L	I	E	A	E		L	s	S	E	Q	I	Н	L	L
	3073/321										/331								
55	TTC CCA				GAG E	ATT	ATT I	GAT D	TTT F		AGA R	CGA R	TTC	L	N N	GGG	L	GAA E	TGT C
	3133/341	N	L	N						3163	/35:	Ĺ							_
	AAC ATC	AAT		CCT			TAT	CAA										TCT	TTG
C 0		N	Λ	P	I	R	Y	Q	R	I			V	F	I	H	A	S	L
60	3193/361 GGC CCT	TTC	TAA	GCT	TAT	GAA	CCT	TGG	ACT	3223 ATA	GGA	CAA	TTG	ACG	GCG	ATT	GAT	TTG	ATC
			N			E				I	G	Q	L	T	Α	I	D	L	I.
	3253/381 AAC AAA		a am	aam	* * 17	mma	* * *	20 20 20	maa	3283	3/39:	L Cmr	amm	a n m	CCT	cee	di di di	C A A	COO
65		E	A	A	N		K			S			L	D	P	G	F	E	L
00	3313/401	-								3343	3/41:	ì							
	CAA TCG																		
	Q S 3373/421		I	L	K	P	I	Q	R	L 3403	3/43:		Y	P	L	L	L	K	E
70	TTA ATC	AAA	ACA	TCA	CCA	GAA	TAT	TCA		CAG	GAC	CCC					TCA		
	L I	K	T	S	P	E	Y	S	K	Q	D	₽		G	S	S	S	S	T
	3433/441 TCA TTC	አልጥ	CND	מיזיים	ጥጥር	GTG	СĊТ	ΔΔΔ	Δ ("Τ"	3463 GCA	3/45: ATG	777 T	GAA	ጥጥር፤	GCD	ΑAΤ	CAA	GTC	ААТ
	S F																		
																	-		



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	3493/46	<u> </u>								352	3/47	1							
	GAG GCG		AGA					ATC	GAA	CAT	TTG	GAA	AAA	CTA	AAA	GAA	AGA	GTA	GGT
	E A		R	R	A	E	N	I	E			E	K	L	ĸ	E	R	V	G
5	3553/48 AAT TGG		aaa	man	ידותת	Time?	ር አጥ	CCM	~ ~ ~		3/49:		מייים א	mmc	CAC	CCA	(T))	COTO	ccc
5	N W	R	G	F	N	L	D	A	Q			L		F	H	G	O	V	G
	3613/50		G	L	14	4.3			¥	_	3/51:		1.3	•		J	v	•	•
	GTT AAA		GCT	GAA	AAT	GAA	AAG	GAA	TAC				CTT	TTT	GAA	AAA	ATC	GTA	TTT
	v K	D	A	E	N	E	ĸ	E	Y	v		Y	L	F	E	к	I	V	F
10	3673/52										3/53:								~
	TTT TTC																		
	F F 3733/54	T	E	I	D	D	T	K	ĸ	S 376	D 3/55:	K 1	Q	E	K	ĸ	S	K	F
	TCG ACA		AAG	AGA	TCA	ACT	TCA	TCA	AAT				TCG	ACT	ACT	AAT	TTG	TTG	GAA
15	S T		K		S	T	s	S	N	L		S	s	T	T	N	L	L	E
	3793/56										3/573								
	TCA ATA																		TAT
	SI	N	N	s	R	K	D	N	T		P		E	L	K	G	R	V	Y
20	3853/58 ATA TCG		מיידי מ	י יי אַ ידי	מממ	ጥጥል	ጥርር	GCA	CCA		3/591 ACT		GGC	тСА	ΑСיπ	СТА	ATC	ATC	TCA
20	I S	E	I	Y	N	I	s		p		T		G	S	T	L	I	I	S
	3913/60	1								3943	3/611	L							
	TGG TCA																		
25	W S	G	R	ĸ	E	S	G	S	F		L 3/631		Y	R	S	E	E	Α	R
23	3973/62: AAC CAA		GAA	DAG	тст	TTA	ССТ	GAT	тта				GAA	ATG	ТАА	ддд	CAA	атт	CAT
	N O		E	K	Ĉ	L	R	D	L			N	E	M	N	ĸ	Q	I	Н
	4033/64										3/651								
20	AAG AAG																		
30	K K 4093/663	L	R.	D	S	D	S	S	F		T 3/671		D	S	A	I	Y	D	Y
	ACG GGT		AGT	ACG	TCA	CCA	GTC	AAT	CAA				CAA	CAA	TAC	TAT	GAT	CAT	CGG
	T G	I	S	T	s	P		N	Q		T			Q	Y	Y	D	H	R
	4153/683										/691								
35	GGC TCT				CGC R			TCA S	TCG		TCC		TTG L	AGT S	ATG M	ATG M	AAG K	AAT N	AAT N
	G S 4213/701	H	S	S	ĸ	п	L1	3	3		3 1/711		ם	3	1-1	1-1	K	14	I
	AGA GTT	AAA	TCT	GGT	GAT	TTG	AGT	AGA	ATA				TCA	ACA	ACA	TTA	GAT	TCT	TTC
	R V	K	S	G	D	L	S	R	I	S	s	T	S	Т	T	L	D	S	\mathbf{F}
40	4273/721	L 	mma	3 3 77	~~~	max.	CCA	20.00	700		731		mom	mma	n ma	mam	max	CATE	aaa
	AGT AAC S N			AAT N	G	S		N	T					L	M	S	S	D	A
	4333/741		_						_	4363	/751	L	_		- "				
	ACC AAA																		
45		-	I	P	T	F	D	V	A	1427		L	L	Y	K	S	T	E	L
	4393/761 TCA GAG		ጥጥር	ልጥጥ	GTC	таа	GCA	CAA	ATT		1/771 TAT		GAC	СТТ	TTA	CAG	AAA	тта	ATC
			L				A	Q	I					L			ĸ	I	I
	4453/781										/791								
50	TCC CAG																		
		_	I	T	ន	N	L	V	A		<i>b</i> /811	V	N	I	S	R	L	R	Y
	4513/801 AAA GAC		GAD	GGA	GAC	ጥጥጥ	GTG	AAT	TTG				CAT	ТДЭ	TGG	GGG	TTA	GTG	CTT
	K D				D					N									L
55	4573/821										/831								
	GAT ATG																		
	D M 4633/841		T	S	E	D	F	Y	Q	T	8	S	N	E	K	R	S	V	T
	GTG TGG		TCT	TGA															
60	V W V																		
		-																	



5	Blast	p line	up o	f S.c. Cdc24p and C.a. Cdc24p	
	S.c.	Cdc24p:	1	MAIQTR-FA	8
10	C.a. (Cdc24p:	1	M R F+ MEHPPAALRTFSTQ	14
10		Cdc24p:		SGTSLSDLKPKPSATSISIPMQNVMNKPVTEQDSLFHICANIRKRLEVLPQLKPFLQI S +SL+ + S+ +S N+ NKP T +D LF+ C +++++L+ +P ++PFL	
	C.a. (Cdc24p:	15	STSSLNSVSTVSSSRIVSSGPVNINNFNKPSTPKDHLFYRCESLKRKLQKIPGMEPFLNG	74
15		-		AYQSSEVLSERQSLLLSQKQHQELLKSNGANRDSSDLAPTLRSSSISTATSLMSMEC A+ +E LSE+Q+L L+Q++ SNG S D A ++ S S S SL M	
		_		AFNQAEQLSEQQALALAQERSNGNGHSNGKRHQSLDGAMNRLSVGSDSSSIQGSLTRMA	
20		-		ISYTNSNPSATPNMEDTLLTFSMGILPITMDCDPVTQLSQLFQQGAPLCILFNSVKPQFF + T+S S PN +TL TF+ G+LP + DP T L +LFQQGAP C+L N + P + NASTSSLISGMPN-NNTLFTFTAGVLPANISVDPATHLWKLFQQGAPFCVLINHILPDSQ	-
		-		LPVIASDDLKVCKKSIYDFILGCKKHFAFNDEELFTISDVFANSTSQLVKVLEVVETLM	
25		-		+PV++SDDL++CKKS+YDF++ K F+DE +FTIS+VF+++ L+K+++V+ L+ IPVVSSDDLRICKKSVYDFLIAVKTQLNFDDENMFTISNVFSDNAQDLIKIIDVINKLL	
				SSPTIFPSKSKTQQIMNAENQHRHQPQQSSKKHNEYVKIIKEFVATERKYVHDLEILDKY	
2.0	C.a. C	Cdc24p:	254	S + + + E K+ +E + TERKYV DLE++ KY EYSDASDSGGGDEDVNMDVQITDERSKVFREIIETERKYVQDLELMCKY	302
30	S.c. C	Cdc24p:	304	RQQLLDSNLITSEELYMLFPNLGDAIDFQRRFLISLEINALVEPSKQRIGALFMH-SKH	362
	C.a. C	Cdc24p:	303	RQ L+++ ++SE++++LFPNL + IDFQRRFL LE N V QRIG++F+H S RQDLIEAENLSSEQIHLLFPNLNEIIDFQRRFLNGLECNINVPIRYQRIGSVFIHASLGE	362
35	S.c. C	dc24p:	363	FKLYEPWSIGQNAAIEFLSSTLHKMRVDESQRFIINNKLELQSFLYKPVQRLCRYPLLVF F YEPW+IGQ AI+ ++ ++ S +++ ELQS++ KP+QRLC+YPLL+F	
	C.a. C	dc24p:.	363	FNAYEPWTIGQLTAIDLINKEAANLKKSSSLLDPGFELQSYILKPIQRLCKYPLLL	
40		_		ELLAESSDDNNTKELEAALDISKNIARSINENQRRTENHQVVKKLYGRV EL+ SS + EL A K +A +NE QRR EN + ++KL RV	•
		_		ELIKTSPEYSKQDPHGSSSSTSFNELLVAKTAMKELANQVNEAQRRAENIEHLEKLKERV	
45		~		VNWKGYRISKFGELLYFDKVFISTTNSSSEPEREFEVYLFEKIIILFSEVVTKKSASSLI NW+G+ + GELL+ +V + +E E+E+ YLFEKI+ F+E+ K + GNWRGFNLDAQGELLFHGQVGVKDAENEKEYVAYLFEKIVFFFTEIDDTKKSDKQE	
	S.c. C	dc24p:	532	LKKKSSTSASISASNITDNNGSPHHSYHKRHSNSSSSNNIHLSSSSAAAIIHSSTNSSDN	591
50	C.a. C	dc24p:	536	K K ST ++SN+ SSS ++ S NS + KKSKFSTRKRSTSSNLSSSTTNLLESINNSRKD	568
50	s.c. C	dc24p:	592	NSNNSSSSSLFKLSANEPKLDLRGRIMIMNLNQIIPQNNRSLNITWESIKEQGNFLLK	
	C.a. C	dc24p:	569	N+ L+L+GR+ I + I N +L I+W KE G+F L+ NTLPLELKGRVYISEIYNISAPNTPGSTLIISWSGRKESGSFTLR	613
55	S.c. C	dc24p:	650	FKNEETRDNWSSCLQQLIHDLKNEQFKARHHSSTSTTSSTAKSSSMMSPTTT+++EE R+ W CL+ L + N+Q + S S+ ++ T S+S ++ +T	701
	C.a. C	dc24p:	614	YRSEEARNQWEKCLRDLKTNEMNKQIHKKLRDSDSSFNTDDSAIYDYTGISTSPVNQSTQ	673
60	S.c. C	dc24p:	702	MNTPNHHNSRQTHDSMASFSSSHMKRVSDVLPKRRTTSSSFESEIKS +H S + H S ++ S RV + TT SF + +	748
00	C.a. C	dc24p:	674	QQYYDHRGSHSSRHHSSSSTLSMMKNNRVKSGDLSRISSTSTTLDSFSNNLNGSPNTTNP	733
	S.c. C	dc24p:	749	ISENFKNSIPESSILFRISYNNNSNNTSSEIFTLLVEKVWNFDDLIMAINSKISN +S + +IP + ++ Y +T SE L+V ++DL+ I S+I SN	
65	C.a. C	dc24p:	734	SLMSSDATKTIPTFDVAIKLLYKSTELSEPLIVNAQIEYNDLLQKIISQIITSN	
	S.c. C	dc24p:	805	THNNNISPITKIKYQDEDGDFVVLGSDEDWNVAKEMLAENNEKFLNIRLY 854 ++++ I++++Y+D++GDFV L SD+DW + +ML + F +	
	C.a. C	dc24p:	788	LVADDVN-ISRLRYKDDEGDFVNLNSDDDWGLVLDMLTSEDFYQTSSNEKRSVTVWVS	844



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FIG. 7A₁

FIG. 7A₂

FIG. 7A

AUG 3 0 2002 PERSONAL STRACEMENT F



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FIG. 7A₁

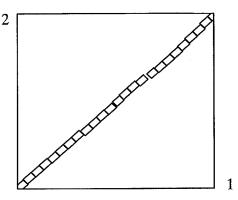
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BLAST 2 SEQUENCES RESULTS VERSION BLASTP 2.0.9 [May-07-1999]

Matrix 0 BLOSUM62	ap open: 11 gap exten	sion: 1
x_dropoff: 50 expect: 10.0	wordsize: 3 <u>Filter</u> [Align

Sequence 1 lcl|S.c. Cdc24p Length 854 (1..854)

Sequence 2 lcl|C.a. Cdc24p Length 844 (1 .. 844)



NOTE: The statistics (bitscore and expect value) is calculated based on the size of nr database

Score = 446 bits (1136), Expect = e-124 Identities = 288/881 (32%), Positives = 464/881 (51%), Gaps = 112/881 (12%)

```
SGTSLSDLKPKPSATSISIPMQNV--MNKPVTEQDSLFHICANIRKRLEVLPQLKPFLQL 66
Query: 9
                                   NKP T +D LF+ C ++++L+ +P ++PFL
          S +SL+ + S+ +S N+
Sbjct: 15 STSSLNSVSTVSSSRIVSSGPVNINNFNKPSTPKDHLFYRCESLKRKLQKIPGMEPFLNQ 74
Query: 67 AYQSSEVLSERQSLLLSQKQHQELLKSNGANRDSSDLAP---TLRSSSISTATSLMSMEG 123
                                                   ++ S S S
          A+ +E LSE+Q+L L+Q++
                                   SNG
                                          SDA
                                                             SL M
Sbjct: 75 AFNQAEQLSEQQALALAQERSNGNGHSNGKRHQSLDGAMNRLSVGSDSSSIQGSLTRMAT 134
Query: 124 ISYTNSNPSATPNMEDTLLTFSMGILPITMDCDPVTQLSQLFQQGAPLCILFNSVKPQFK 183
           + T+S S PN +TL TF+ G+LP + DP T L +LFQQGAP C+L N + P
Sbjct: 135 NASTSSLISGMPN-NNTLFTFTAGVLPANISVDPATHLWKLFQQGAPFCVLINHILPDSQ 193
Query: 184 LPVIASDDLKVCKKSIYDFILGCKKHFAFNDEELFTISDVFANSTSQLVKVLEVVETLMN 243
           +PV++SDDL++CKKS+YDF++ K F+DE +FTIS+VF+++ L+K+++V+ L+
Sbjct: 194 IPVVSSDDLRICKKSVYDFLIAVKTQLNFDDENMFTISNVFSDNAQDLIKIIDVINKLLA 253
Query: 244 SSPTIFPSKSKTQQIMNAENQHRHQPQQSSKKHNEYVKIIKEFVATERKYVHDLEILDKY 303
                                       + +E K+ +E + TERKYV DLE++ KY
                 S
                      + +
Sbjct: 254 EYSDASDSGGGDEDV------NMDVQITDERSKVFREIIETERKYVQDLELMCKY 302
Query: 304 RQQLLDSNLITSEELYMLFPNLGDAIDFQRRFLISLEINALVEPSKQRIGALFMH-SKHF 362
          RQ L+++ ++SE+++LFPNL + IDFQRRFL LE N V
                                                       QRIG++F+H S
Sbjct: 303 RQDLIEAENLSSEQIHLLFPNLNEIIDFQRRFLNGLECNINVPIRYQRIGSVFIHASLGP 362
Query: 363 FKLYEPWSIGQNAAIEFLSSTLHKMRVDESQRFIINNKLELQSFLYKPVQRLCRYPLLVK 422
          F YEPW+IGQ AI+ ++ ++ S +++ ELQS++ KP+QRLC+YPLL+K
Sbjct: 363 FNAYEPWTIGQLTAIDLINKEAANLKKSSS---LLDPGFELQSYILKPIQRLCKYPLLLK 419
```



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```
Query: 551 --NGSPHHSYHKRHSNSSSSNNIHLS-----SSSAAAIIHSSTNSSDNNSNNSSSS 599
            GPH
                     + S+ + +S
                                             + I S+ ++ N N SSS
Sbjct: 477 FWRGDPQHESFILKLRNEESHKLWMSVLNRLLWKNEHGSPKDIRSAASTPANPVYNRSSS 536
Ouery: 600 SLFKLSANEPKLD-LRGRIMIMNLN---QIIPQNNRSLNITWESIKEQGNFLLKFKNEET 655
                   D LR + N+N I +++S T + K+
Sbjct: 537 QTSK-GYNSSDYDLLRTHSLDENVNSPTSISSPSSKSSPFTKTTSKDT-----KSATT 588
Query: 656 RDNWSSCLQQLIHDLKNEQFKARHHSSTST----TSSTAKSSSMMSPTTTMNT--PNHH 708
                  +L +
                           R
                                  +TST
                                           +SSTA
                                                  `S +S + +N+
Sbjct: 589 TDERPSDFIRLNSEESVGTSSLRTSQTTSTIVSNDSSSTASIPSQISRISQVNSLLNDYN 648
Query: 709 NSRQTH------DSMASF---SSSHMKRVSD------VLPKRRTTSSSFESE 745
                           S++ F SSS +++ D
                                                    + P++ + S+ +S+
Sbjct: 649 YNRQSHITRVYSGTDDGSSVSIFEDTSSSTKQKIFDQPTTNDCDVMRPRQYSYSAGMKSD 708
Query: 746 IKSISENFKNSIPESSILFRISYNNNSNNTSSSEI----FTLLVEKVWNFDDLIMAINSK 801
                   S+ SS +S N +N + L+V
                                                       FD+L+ + K
Sbjct: 709 GSLLPSTKHTSLSSSSTSTSLSVRNTTNVKIRLRLHEVSLVLVVAHDITFDELLAKVEHK 768
Query: 802 IS--NTHNNNISPITKIKYQDEDGDFVVLGSDEDWNVAKE 839
              + ++KY DEDGDF+ + SDED +A E
Sbjct: 769 IKLCGILKQAVPFRVRLKYVDEDGDFITITSDEDVLMAFE 808
CPU time: 0.26 user secs. 0.04 sys. secs 0.30 total secs.
Gapped
Lambda
          0.0470
                   0.230
  0.270
Matrix: BLOSUM62
Gap Penalties: Existence: 11, Extension: 1
Number of Hits to DB: 10384
Number of Sequences: 0
Number of extensions: 671
Number of successful extensions: 13
Number of sequences better than 10.0: 1
Number of HSP's better than 10.0 without gapping: 1
Number of HSP's successfully gapped in prelim test: 0
Number of HSP's that attempted gapping in prelim test: 0
Number of HSP's gapped (non-prelim): 2
length of query: 834
length of database: 90,077,593
effective HSP length: 61
effective length of query: 773
effective length of database: 83489227
effective search space: 64537172471
effective search space used: 64537172471
T: 9
A: 40
X1: 16 ( 7.3 bits)
x2: 128 (49.9 bits)
X3: 128 (49.9 bits)
S1: 41 (21.7 bits)
S2: 73 (32.8 bits)
```



LEOTIDE SEQUENCES AND PROTEIN SI ENCE.

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Appl. No.: 10/054,399 Atty Docket: DYOU13.1A2CP1

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FIG. 7B₁

FIG. 7B₂

FIG. 7B



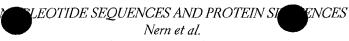


FIG. 7B₁

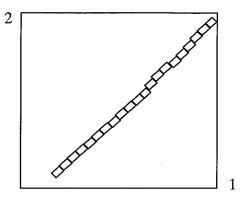
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BLAST 2 SEQUENCES RESULTS VERSION BLASTP 2.0.9 [May-07-1999]

Matrix 0	BLOSUM6	2 •	gap open:	gap ext	ension: 1
x_dropoff:	50 expect:	10.0 w	ordsize: 3	Filter	Align

Sequence 1 lcl|S.c. Cdc24p Length 854 (1..854)

Sequence 2 lcl|S.p. Cdc24p Length 834 (1..834)



NOTE: The statistics (bitscore and expect value) is calculated based on the size of nr database

Score = 238 bits (601), Expect = 1e-61 Identities = 212/760 (27%), Positives = 348/760 (44%), Gaps = 94/760 (12%)

```
Query: 156 DPVTQLSQLFQQGAPLCILFNSVKPQFKLPVIASDDLK---VCKKSIYDFILGCKKHFAF 212
                  + G PLC LFN + + KL V +S L+ VCK S+Y F+L CK
Sbjct: 67 DPVTEIWLFCRLGYPLCALFNCLPVKQKLEVNSSVSLENTNVCKASLYRFMLMCKNELGL 126
Query: 213 NDEELFTISDVFANSTSQLVKVLEVVETLMNSSPTIFPSKSKTQQIMNAENQHRHQPQQS 272
           D LF+IS+++ ST+ LV+ L+ +E L+
                                              +KS +
                                                      + ++
Sbjct: 127 TDAALFSISEIYKPSTAPLVRALQTIELLLKKYEVSNTTKSSSTPSPSTDDNVPTGTLNS 186
Query: 273 SKKHNEYVKIIKEFVATERKYVHDLEILDKYRQQLLDSNLITSEELYMLFPNLGDAIDFQ 332
                  ++ E
                        TE KY+ DLE L Y L +++ + +F NL + +DFQ
Sbjct: 187 LIASGR--RVTAELYETELKYIQDLEYLSNYMVILQQKQILSQDTILSIFTNLNEILDFQ 244
Query: 333 RRFLISLEINALVEPSKQRIGALFMHSKHFFKLYEPWSIG-QNAAIEFLSSTLHKMRVDE 391
          RRFL+ LE+N + +OR+GALF+ + F +Y+ + NA + +
Sbjct: 245 RRFLVGLEMNLSLPVEEQRLGALFIALEEGFSVYQVFCTNFPNAQQLIIDNQNQLLKVAN 304
Query: 392 SQRFIINNKLELQSFLYKFVQRLCRYPLLVKELL-AESSDDNNTKELEAALDISKNIARS 450
                   EL + L KP+QR+C+YPLL+ +LL S
                                                    +EL+ +
Sbjct: 305 ----LLEPSYELPALLIKPIQRICKYPLLLNQLLKGTPSGYQYEEELKQGMACVVRVANQ 360
Query: 451 INENQRRTENHQVVKKLYGRVVNWKGYRISKFGELLYFDKVFISTTNSSSEPEREFEVYL 510
          +NE +R EN + +L RV++WKGY + FG+LL +D V +
                                                         ++ ERE+ VYL
Sbjct: 361 VNETRRIHENRNAIIELEQRVIDWKGYSLQYFGQLLVWDVVNV----CKADIEREYHVYL 416
Query: 511 FEKIIILFSEVVT-KKSASSLILKKKSSTSAS----ISASNITDN----- 550
                                     S
          FEKI++
                  E+ T K+ A S+ + KK+
                                              I SNIT
Sbjct: 417 FEKILLCCKEMSTLKRQARSISMNKKTKRLDSLQLKGRILTSNITTVVPNHHMGSYAIQI 476
```

FIG. 7B₂

```
Query: 423 ELLAE-----SSDDNNTKELEAALDISKNIARSINENQRRTENHQVVKKLYGRV 471
          EL+ SS + EL A K +A +NE QRR EN + ++KL RV
Sbjct: 420 ELIKTSPEYSKQDPHGSSSSTSPNELLVAKTAMKELANQVNEAQRRAENIEHLEKLKERV 479
Query: 472 VNWKGYRISKFGELLYFDKVFISTTNSSSEPEREFEVYLFEKIIILFSEVVTKKSASSLI 531
           NW+G+ + GELL+ +V + +E E+E+ YLFEKI+ F+E+ K +
Sbjct: 480 GNWRGFNLDAQGELLFHGQVGV----KDAENEKEYVAYLFEKIVFFFTEIDDTKKSDKQE 535
Query: 532 LKKKSSTSASISASNITDNNGSPHHSYHKRHSNSSSSNNIHLSSSSAAAIIHSSTNSSDN 591
           K K ST ++SN+
                                                   SSS ++ S NS
Sbjct: 536 KKSKFSTRKRSTSSNL-----
                                               ----SSSTINLLESINNSRKD 568
Query: 592 NSNNSSSSSLFKLSANEPKLDLRGRIMIMNLNQIIPQN--NRSLNITWESIKEQGNFLLK 649
                            L+L+GR+ I + I N
                                                  +L I+W
                                                          KE G+F L+
                 -----LPLELKGRVYISEIYNISAPNTPGSTLIISWSGRKESGSFTLR 613
Sbjct: 569 NT--
Query: 650 FKNEETRDNWSSCLQQLIHDLKNEQFKARHHSSTSTTSS-----TAKSSSMMSPTTT 701
          +++EE R+ W CL+ L + N+Q + S S+ ++ T S+S ++ +T
Sbjct: 614 YRSEEARNQWEKCLRDLKTNEMNKQIHKKLRDSDSSFNTDDSAIYDYTGISTSPVNQSTQ 673
Query: 702 MNTPNHHNSRQT--HDSMASFSSSHMKRVS----DVLPKRRTTSSSFESEIKS----- 748
              +H S + H S ++ S RV + TT SF + +
Sbjct: 674 QQYYDHRGSHSSRHHSSSSTLSMMKMNRVKSGDLSRISSTSTTLDSFSNNLNGSPNTTNP 733
Query: 749 -- ISENFKNSIPESSILFRISYNNNSNNTSSEIFTLLVEKVWNFDDLIMAINSKI--SN 804
            +S + +IP + ++ Y +T SE L+V ++DL+ I S+I SN
Sbjct: 734 SLMSSDATKTIPTFDVAIKLLY----KSTELSE--PLIVNAQIEYNDLLQKIISQIITSN 787
Query: 805 THNNNISPITKIKYQDEDGDFVVLGSDEDWNVAKEMLAENN 845
             ++++ I++++Y+D++GDFV L SD+DW + +ML
Sbjct: 788 LVADDVN-ISRLRYKDDEGDFVNLNSDDDWGLVLDMLTSED 827
                                0.02 sys. secs
                                                        0.28 total secs.
CPU time:
           0.26 user secs.
Gapped
Lambda
          ĸ
          0.0470
   0.270
                    0.230
Matrix: BLOSUM62
Gap Penalties: Existence: 11, Extension: 1
Number of Hits to DB: 12253
Number of Sequences: 0
Number of extensions: 709
Number of successful extensions: 15
Number of sequences better than 10.0: 1
Number of HSP's better than 10.0 without gapping: 1
Number of HSP's successfully gapped in prelim test: 0
Number of HSP's that attempted gapping in prelim test: 0
Number of HSP's gapped (non-prelim): 1
length of query: 844
length of database: 90,077,593
effective HSP length: 63
effective length of query: 781
effective length of database: 83353792
effective search space: 65099311552
effective search space used: 65099311552
T: 9
A: 40
X1: 16 ( 7.2 bits)
X2: 128 (49.9 bits)
X3: 128 (49.9 bits)
S1: 42 (21.9 bits)
$2: 73 (32.8 bits)
```



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Sc KLPVIASDDLKVCKKSIYDFIL (SEQ ID No 25) ++PV++SDDL++CKKS+YDF++ Ca QIPVVSSDDLRICKKSVYDFLI (SEQ ID No 26)

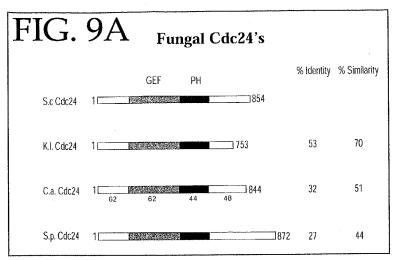
 $Sc = Saccharomyces\ cerevisiae$

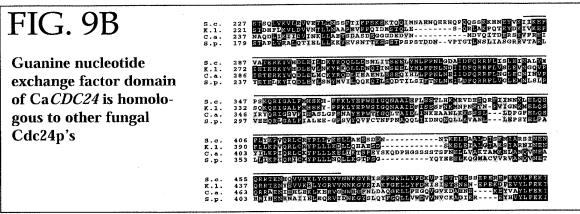
Ca = Candida albicans

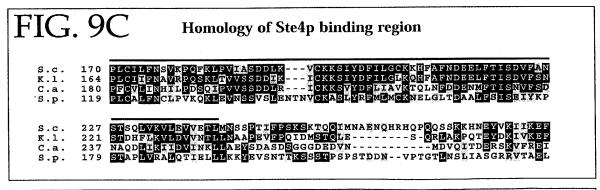
FIG. 8



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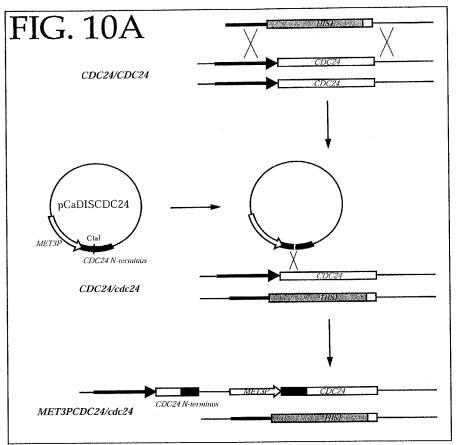


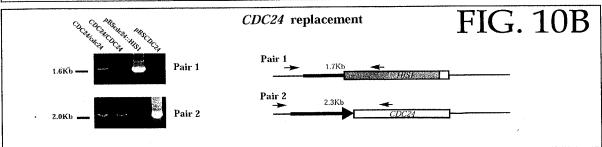
08/30/02

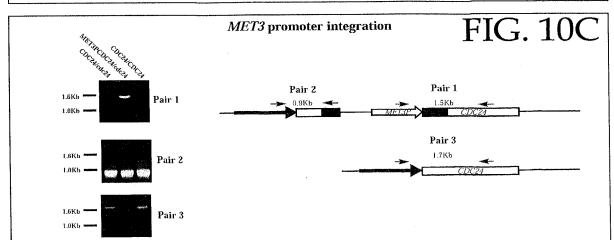
N FOTIDE SEQUENCES AND PROTEIN UENCES

Nern et al.

Appl. No.: 10/054,399 Atty Docket: DYOU13.1A2CP1

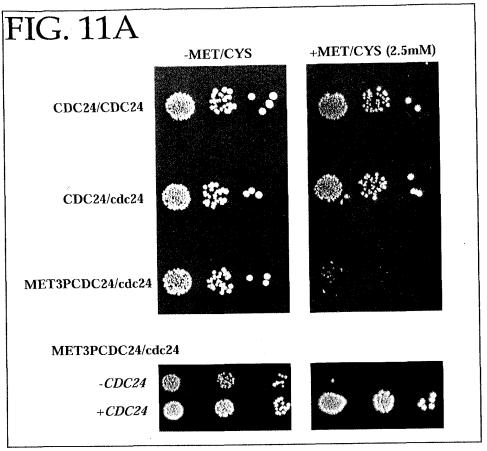


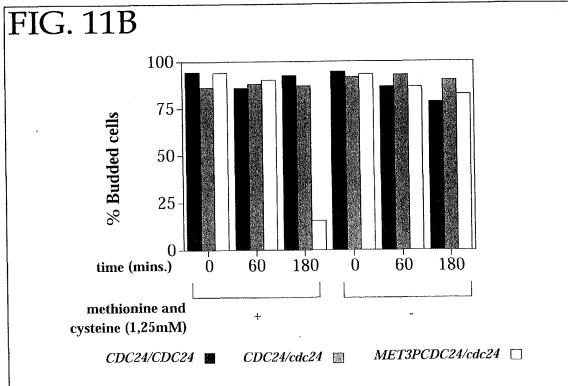






N EOTIDE SEQUENCES AND PROTEIN UENCES
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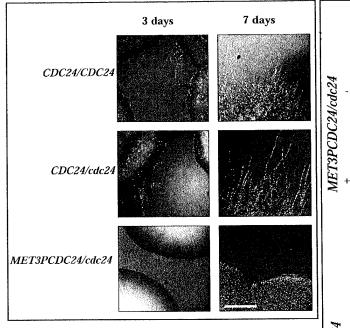


N' L'EOTIDE SEQUENCES AND PROTEIN VIENCES Nern et al. Appl. No.: 10/054,399 Atty Docket: DYOU13.1A2CP1

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FIG. 12A

FIG. 12B



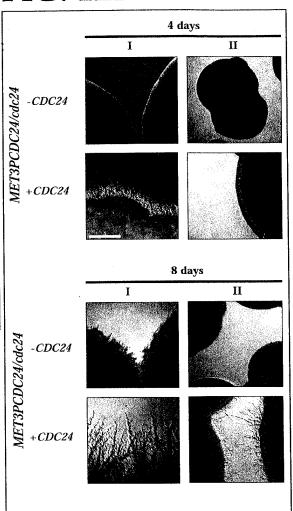
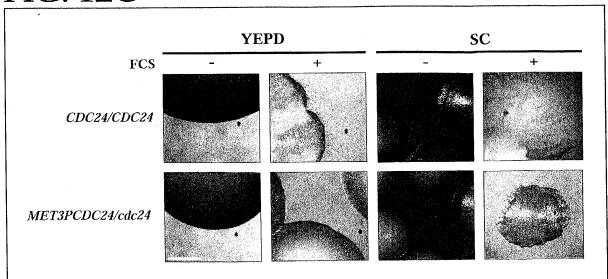


FIG. 12C





N' FOTIDE SEQUENCES AND PROTEIN S VENCES Nem et al. Appl. No.: 10/054,399 Atty Docket: DYOU13.1A2CP1

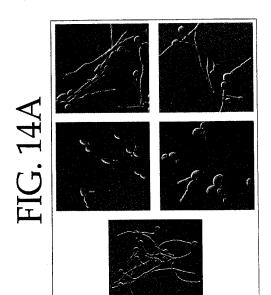


FIG. 14B

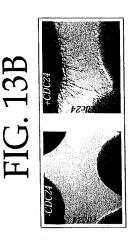
FIG. 13A

with type

cate 24

cate 24

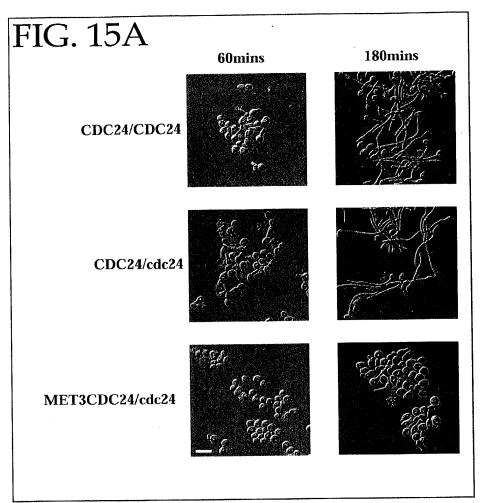
but it

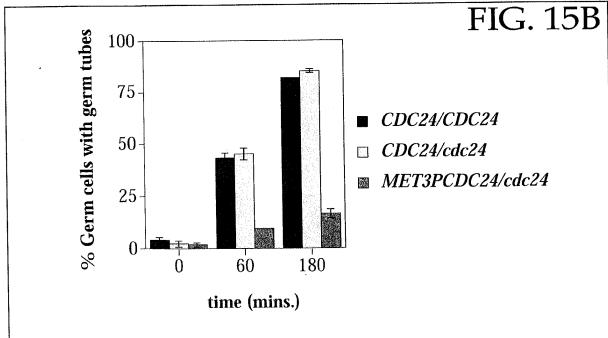


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EOTIDE SEQUENCES AND PROTEIN **UENCES** Nern et al.

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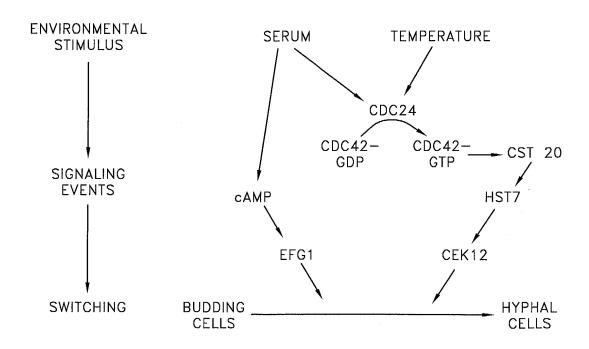


FIG. 16



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FIG. 17_A

FIG. 17_B

FIG. 17_C

FIG. 17_D

FIG. 17_E

FIG. 17



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A. *cdc24* (wt)

5

SEQ. I.D. NO:1

DNA:

cccctctgtatactttteaactctgtgaageegeaatttaaattaceggtaatageatctgacgatttgaaagtctgtaaaaaatccatttatgactt
tatattgggctgcaagaaacactttgcatttaacgatgaggagettttcactatatccgacgtttttgccaactcgacgtcccagctggtcaaagt
gctagaagtagtagaaacgctaatgaattccagc

SEQ. I.D. NO:2

Protein:

15 PLCILFNSVKPQFKLPVIASDDLKVCKKSIYDFILGCKKHFAFNDEELFTISDVFANSTSQ LVKVLEVVETLMNSS

B. cdc24-m1

20 SEQ. I.D. NO:3

DNA:

ccctctgtatacttttcaactctgtgaagccgcaatttaaattaccggtaatagcatttgacgatttgaaagtctgtaaaaaatccatttatgacttt atattgggctgcaagaaacactttgcatttaacgatgaggagcttttcactatatccgacgtttttgccaactcgacgtcccagctggtcaaagt gctagaagtagtagaaacgctaatgaattccagc

25

SEQ. I.D. NO:4

Protein:

PLCILFNSVKPQFKLPVIAFDDLKVCKKSIYDFILGCKKHFAFNDEELFTISDVFANSTSQ LVKVLEVVETLMNSS

30



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FIG. $17_{\rm B}$

C. cdc24-m2

5 SEQ. I.D. NO:5

DNA:

cccctctgtatacttttcaactctgtgaagccgcaatttaaattaccggtaatagcatctggcgatttgaaagtctgtaaaaaatccatttatgactt tatattgggctgcaagaaacactttgcatttaacgatgaggagcttttcactatatccgacgtttttgccaactcgacgtcccagctggtcaaagt gctagaagtagtagaaacgctaatgaattccagc

10

SEQ. I.D. NO:6

Protein:

PLCILFNSVKPQFKLPVIASGDLKVCKKSIYDFILGCKKHFAFNDEELFTISDVFANSTSQ LVKVLEVVETLMNSS

15

D. cdc24-m3

SEQ. I.D. NO:7

DNA:

20 cccctctgtatacttttcaactctgtgaagccgcaatttaaattaccggtaatagcacctgacgatttgaaagtctgtaaaaaatccatttatgactt tatattgggctgcaagaaacactttgcatttaacgatgaggagcttttcactatatccgacgtttttgccaactcgacgtcccagctggtcaaagt gctagaagtagtagaaacgctaatgaattccagc

SEQ. I.D. NO:8

25 Protein:

PLCILFNSVKPQFKLPVIAPDDLKVCKKSIYDFILGCKKHFAFNDEELFTISDVFANSTSQ LVKVLEVVETLMNSS



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FIG. 17_C

SEQ ID NO. 10

STE4 DNA sequence (wild-type)

ATGGCACATCAGATGGACTCGATAACGTATTCTAATAATGTCACCCAACAGTATATACAACCACAAAGTCTACAGGA ATAAAGCAAAACACAAGATACAAGATGCAAGCTTATTCCAGATGGCCAACAAGTTACTTCGTTGACCAAAAATAAGATC AACTTAAAGCCAAATATCGTGTTGAAAGGCCATAATAATAAAATCTCAGATTTTCGGTGGAGTCGAGATTCAAAACGTAT TTTGAGTGCAAGTCAAGATGGCTTTATGCTTATATGGGACAGTGCTTCAGGTTTAAAACAGAACGCTATTCCATTAGATT CTCAATGGGTTCTTTCCTGCGCTATTTCGCCATCGAGTACTTTGGTAGCAAGCGCAGGATTAAACAATAACTGTACCATT TATAGAGTTTCGAAAGAAAACAGAGTAGCGCAAAACGTTGCGTCAATTTTCAAAGGACATACTTGCTATATTTCTGACAT TGAATTTACAGATAACGCACATATATTGACAGCAAGTGGGGATATGACATGTGCCTTGTGGGATATACCGAAAGCAAAGA GGGTGAGAGAATATTCTGACCATTTAGGTGATGTTTTGGCATTAGCTATTCCTGAAGAGCCAAACTTAGAAAATTCTTCG 10 CGTTAACGATAGTGATATTAATGCACTTCGTTTTTTCAAAGACGGGATGTCGATTGTTGCAGGAAGTGACAATGGTGCGA TAAATATGTATGATTTAAGGTCGGACTGTTCTATTGCTACTTTTTCTCTTTTTCGAGGTTATGAAGAACGTACCCCTACC CCTACTTATATGGCAGCTAACATGGAGTACAATACCGCGCAATCGCCACAAACTTTAAAATCAACAAGCTCAAGCTATCT ${\tt AGACAACCAAGGCGTTGTTTCTTTAGATTTTAGTGCATCTGGAAGATTGATGTACTCATGCTATACAGACATTGGTTGTG}$ 15 TTGTGTGGGATGTATTAAAAGGAGAGATTGTTGGAAAATTAGAAGGTCATGGTGGCAGAGTCACTGGTGTGCGCTCGAGT CCAGATGGGTTAGCTGTATGTACAGGTTCATGGGACTCAACCATGAAAATATGGTCTCCAGGTTATCAATAG

SEQ ID NO. 11 20 Ste4 Protein sequence (wild-type)

MAHQMDSITYSNNVTQQYIQPQSLQDISAVEEEIQNKIEAARQESKQLHAQINKAKHKIQDASLFQMANKVTSLTKNKIN LKPNIVLKGHNNKISDFRWSRDSKRILSASODGFMLIWDSASGLKQNAIPLDSQWVLSCAISPSSTLVASAGLNNNCTIY ${\tt RVSKENRVAQNVASIFKGHTCYISDIEFTDNAHILTASGDMTCALWDIPKAKRVREYSDHLGDVLALAIPEEPNLENSSN}$ TFASCGSDGYTYIWDSRSPSAVQSFYVNDSDINALRFFKDGMSIVAGSDNGAINMYDLRSDCSIATFSLFRGYEERTPTP TYMAANMEYNTAQSPQTLKSTSSSYLDNQGAVSLDFSASGRLMYSCYTDIGCVVWDVLKGEIVGKLEGHGGRVTGVRSSP DGLAVCTGSWDSTMKIWSPGYQ

30

25

SEQ ID NO. 12 ste4-o15 DNA sequence (mutant)

ATGGCACATCAGATGGACTCGATAACGTATTCTAATAATGTCACCCAACAGTATATACAACCACAAAGTCTACAGGA 35 TATCTCTGCAGTGGAGGAGAAATTCAAAATAAAATAGAGGCCGCCAGACAAGAGAGTAAACAGCTTCATGCTCAAATAA ATAAAGCAAAACACAAGATACAAGATGCAAGCTTATTCCAGATGGCCAACAAAGTTACTTCGTTGACCAAAAATAAGATC AACTTAAAGCCAAATATCGTGTTGAAAGGCCATAATAATAAAATCTCAGATTTTCGGTGGAGTCGAGATTCAAAACGTAT



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FIG. 17_D

TTTGAGTGCAAGTCAAGATGGCTTTATGCTTATATGGGACAGTGCTTCAGGTTTAAAACAGAACGCTATTCCATTAGATT $\tt CTCAATGGGTTCTTTCCTGCGCTATTTCGCCATCGAGTACTTTGGTAGCAAGCGCAGGATTAAACAATAACTGTACCATT$ ${\tt TATAGAGTTTCGAAAGAAAACAGAGTAGCGCAAAACGTTGCGTCAATTTTCAAAGGACATACTTGCTATATTTCTGACAT}$ TGAATTTACAGATAACGCACATATATTGACAGCAAGTGGGGATATGACATGTGCCTTGTGGGATATACCGAAAGCAAAGA GGGTGAGAGAATATTCTGACCATTTAGGTGATGTTTTGGCATTAGCTATTCCTGAAGAGCCAAACTTAGAAAATTCTTCG CGTTAACGATAGTGATATTAATGCACTTCGTTTTTTCAAAGACGGGATGTCGATTGTTGCAGGAAGTGACAATGGTGCGA TAAATATGTATGATTTAAGGTCGGACTGTTCTATTGCTACTTTTTCTCTTTTTCGAGGTTATGAAGAACGTACCCCTACC $\tt CCTACTTATATGGCAGCTAACATGGAGTACAATACCGCGCAATCGCCACAAACTTTAAAATCAACAAGCTCAAGCTATCT$ 10 AGACAACCAAGGCGTTGTTTCTTTAGATTTTAGTGCATCTGGAAGATTGATGTACTCATGCTATACAGACATTGGTTGTG TTGTGTGGGATGTATTAAAAGGAGAGTTGTTGGAAAATTAGAAGGTCATGGTGGCAGAGTCACTGGTGTGCGCTCGAGT CCAGATGGGTTAGCTGTATGTACAGGTTCATGGGACTCAACCATGAAAATATGGTCTCCAGGTTATCAATAG

SEO ID NO. 13

15 Ste4-o15 Protein sequence (mutant)

MAHQMDSITYSNNVTQQYIQPQSLQDISAVEEEIQNKIEAARQESKQLHAQINKAKHKIQDASLFQMANKVTSLTKNKIN LKPNIVLKGHNNKISDFRWSRDSKRILSASQDGFMLIWDSASGLKQNAIPLDSQWVLSCAISPSSTLVASAGLNNNCTIY RVSKENRVAQNVASIFKGHTCYISDIEFTDNAHILTASGDMTCALWDIPKAKRVREYSDHLGDVLALAIPEEPNLENSSN TFASCGSDGYTYIWDSRSPSAVOSFYVNDSDINALRFFKDGMSIVAGSDNGAINMYDLRSDCSIATFSLFRGYEERTPTP TYMAANMEYNTAOSPOTLKSTSSSYLDNOGAVSLDFSASGRLMYSCYTDIGCVVWDVLKGEIVGKLEGHGGRVTGVRSSP DGLAVCTGSWDSTMKIWSPGYQ

SEO ID NO. 14

20

30

35

25 ste4-o17 DNA sequence (mutant)

ATGGCACATCAGATGGACTCGATAACGTATTCTAATAATGTCACCCAACAGTATATACAACCACAAAGTCTACAGGA TATCTCTGCAGTGGAGGAAGAATTCAAAATAAAATAGAGGCCGCCAGACAAGAGAGTAAACAGCTTCATGCTCAAATAA ATAAAGCAAAACACAAGATACAAGATGCAAGCTTATTCCAGATGGCCAACAAAGTTACTTCGTTGACCAAAAATAAGATC AACTTAAAGCCAAATATCGTGTTGAAAGGCCATAATAATAAAATCTCAGATTTTCGGTGGAGTCGAGATTCAAAACGTAT TTTGAGTGCAAGTCAAGATGGCTTTATGCTTATATGGGACAGTGCTTCAGGTTTAAAACAGAACGCTATTCCATTAGATT CTCAATGGGTTCTTTCCTGCGCTATTTCGCCATCGAGTACTTTGGTAGCAAGCGCAGGATTAAACAATAACTGTACCATT TATAGAGTTTCGAAAGAAACAGAGTAGCGCAAAACGTTGCGTCAATTTTCAAAGGACATACTTGCTATATTTCTGACAT TGAATTTACAGATAACGCACATATATTGACAGCAAGTGGGGATATGACATGTGCCTTGTGGGATATACCGAAAGCAAAGA GGGTGAGAGATATTCTGACCATTTAGGTGATGTTTTGGCATTAGCTATTCCTGAAGAGCCAAACTTAGAAAATTCTTCG CGTTAACGATAGTGATATTAATGCACTTCGTTTTTTCAAAGACGGGATGTCGATTGTTGCAGGAAGTGACAATGGTGCGA



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FIG. 17_E

TAAATATGTATGATTTAAGGTCGGACTGTTCTATTGCTACTTTTTCTCTTTTTCGAGGTTATGAAGAACGTACCCCTACC
CCTACTTATATGGCAGCTAACATGGAGTACAATACCGCGCAATCGCCACAAACTTTAAAATCAACAAGCTCAAGCTATCT
AGACAACCAAGGCGTTGTTTCTTTAGATTTTAGTGCATCTGGAAGATTGATGTACTCATGCTATACAGACATTGGTTGTG
TTGTGTGGGATGTATTAAAAGGAGAGATTGTTGGAAAAATTAGAAGGTCATGGTGGCAGGTCACTGAGT
CCAGATGGGTTAGCTGTATGTACAGGTTCATGGGACTCAACCATGAAAATATGGTCTCCAGGTTATCAATAG

SEQ ID NO. 15

Ste4-o17 Protein sequence (mutant)

10 MAHQMDSITYSNNVTQQYIQPQSLQDISAVEEEIQNKIEAARQESKQLHAQINKAKHKIQDASLFQMANKVTSLTKNKIN
LKPNIVLKGHNNKISDFRWSRDSKRILSASQDGFMLIWDSASGLKQNAIPLDSQWVLSCAISPSSTLVASAGLNNNCTIY
RVSKENRVAQNVASIFKGHTCYISDIEFTDNAHILTASGDMTCALWDIPKAKRVREYSDHLGDVLALAIPEEPNLENSSN
TFASCGSDGYTYIWDSRSPSAVQSFYVNDSDINALRFFKDGMSIVAGSDNGAINMYDLRSDCSIATFSLFRGYEERTPTP
TYMAANMEYNTAQSPQTLKSTSSSYLDNQGAVSLDFSASGRLMYSCYTDIGCVVWDVLKGEIVGKLEGHGGRVTGVRSSP
DGLAVCTGSWDSTMKIWSPGYQ

SEQ I.D. No:16 refers to the epitope sequence "Tyr Pro Tyr Asp Val Pro Asp Tyr Ala".

SEQ I.D. No:17 refers to TEV protease recognition sequence "Gln Asn Leu Tyr Phe Gln Gly".

SEQ I.D. No:18 refers to peptide sequence "QFKLPVIAFDDLKVCKKSI".

SEQ I.D. No:19 refers to peptide sequence "QFKLPVIASGDLKVCKKSI".

SEQ I.D. No: 20 refers to peptide sequence "QFKLPVIAPDDLKVCKKSI".

SEQ I.D. No:21 refers to peptide sequence "QFKLPVIASDDLKVCKKSI".

SEQ I.D. No:22 refers to peptide sequence "QYEFDVILSPELKVQMKTI".